Serial#: 10/520,078 STRUCTURE SEARCH

=> FILE HCAPLUS

FILE 'HCAPLUS' ENTERED AT 11:58:56 ON 30 JUN 2009
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FILE COVERS 1907 - 30 Jun 2009 VOL 151 ISS 1

FILE LAST UPDATED: 29 Jun 2009 (20090629/ED)

REVISED CLASS FIELDS (/NCL) LAST RELOADED: Apr 2009

USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Apr 2009

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the second quarter of 2009.

CAS Information Use Policies apply and are available at:

http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.
'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

=> D STAT QUE L12

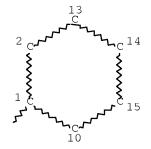
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017 S18

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Page 1-A



Page 1-B



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GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L4 (47768)SEA FILE=REGISTRY SSS FUL L3

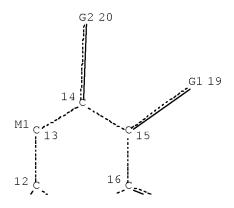
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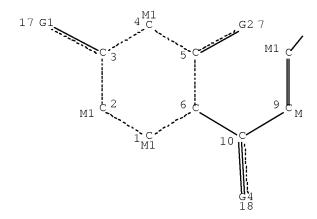
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X 22 Ak 23H 25 O M1 S 26

Page 1-A



Page 1-B



Page 2-A



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Page 2-B VAR G1=22/23/24/25/26

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VAR G2=27/28/29/30
 VAR G4=31/32/33
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 NUMBER OF NODES IS 33
 STEREO ATTRIBUTES: NONE
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^{=&}gt; D L12 IBIB ABS HITSTR 1-5

L12 ANSWER 1 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2009:258682 HCAPLUS Full-text

DOCUMENT NUMBER: 150:306643

TITLE: Preparation of diphenylheteroaryl and chalcone

derivatives as PPAR agonists

INVENTOR(S): Hibbs, David Edward; Salam, Noeris Kris; Roubin,

Rebecca; Matin, Azadeh; Gavande, Navnath S.

PATENT ASSIGNEE(S): The University of Sydney, Australia

SOURCE: PCT Int. Appl., 67pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT	NO.			KIN	D	DATE			APPL	ICAT	ION :	NO.		D.	ATE	
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	FI,	GB,	GD,	GE,	GH,	GM,	GT,	HN,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,
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	ME,	MG,	MK,	MN,	MW,	MX,	MY,	MZ,	NA,	NG,	NI,	NO,	NZ,	OM,	PG,	PH,
	PL,	PT,	RO,	RS,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SM,	ST,	SV,	SY,	ТJ,
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	IE,	IS,	ΙT,	LT,	LU,	LV,	MC,	MT,	NL,	NO,	PL,	PT,	RO,	SE,	SI,	SK,
	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,
	TG,	BW,	GH,	GM,	KΕ,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,
	AM,	ΑZ,	BY,	KG,	KΖ,	MD,	RU,	ТJ,	TM							
PRIORITY APE	.:						AU 2	007-	9046	74		A 2	0070	829		
OTHER SOURCE		MAR:	PAT	150:	3066	43										
GI																

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Title compds. I [A = heteroaryl ring (optionally substituted with halo, alkyl, haloalkyl, etc.); R1-R10 = H, hydroxy, halo, etc.; or their pharmaceutically acceptable salts] and II [L = alkylene or alkenylene; R11-R15 = H, hydroxyl, halo, etc.; R16-R20 = H, hydroxyl, halo, etc.; or their pharmaceutically acceptable salts] were prepared For example, reaction of resorcinol with 4-fluorophenylacetic acid in BF3·OEt2 followed by cyclocondensation with acetic anhydride and treatment with NH2NH2·H2O afforded compound III, which showed PPAR- γ fold activation activity (5.3 at 25 μ M) compared to rosiglitazone (4 at 25 μ M). Compds. I and II are claimed useful for the treatment of type II diabetes, obesity, etc.

IT 961-29-5P 220430-82-0P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of diphenylheteroaryl and chalcone derivs. as PPAR agonists for treatment of type II diabetes, obesity, etc.)

RN 961-29-5 HCAPLUS

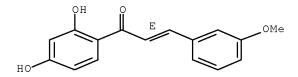
CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

RN 220430-82-0 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(3-methoxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.



REFERENCE COUNT: 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2007:1470011 HCAPLUS Full-text

DOCUMENT NUMBER: 148:100385

TITLE: Preparation of 1,3-diphenylpropane derivatives,

particularly 2-[4-(3-oxo-3-phenylpropyl)phenoxy]-2-methylpropanoic acids and related derivatives, as PPAR agonists for treating diseases especially dyslipidemia Delhomel, Jean-Francois; Hanf, Remy; Caumont-Bertrand,

PATENT ASSIGNEE(S): Genfit, Fr.

SOURCE: PCT Int. Appl., 108pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PAT	TENT	NO.			KIN	D	DATE		ì	APPL	ICAT	ION I	NO.		D	ATE	
WO	 2007	1478	 79		A1	_	 2007	1227		MO 2	 007-:	EP56.	 224		2	0070	 621
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AU	2007	2629	38		A1		2007	1227		AU	2007-	2629	38		2	0070	621
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EP	2046	715			A1		2009	0415		EΡ	2007-	7302	96		2	0070	621
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KR	2009	0355	35		Α		2009	0409		KR	2009-	7013	19		2	0090	121
PRIORITY	APP	LN.	INFO	.:						FR	2006-	5540			A 2	0060	621
										WO	2007-	EP56	224	1	W 2	0070	621
OTHER SC	DURCE	(S):			MAR	PAT	148:	1003	85								

GΙ

AΒ Title compds. I [X1 = R1, G1R1; X2 = halo, R2, G2R2; X3 = R3, G3R3; X4 = halo, R4, G4R4; X5 = R5, G5R5; R1 = H, nonhalogenated alkyl; R2 = H, alkyl; R3-R5 =independently H, (un) substituted alkyl; G1-G5 = independently O, S; with at least one of X3-X5 = R3, G3R3, R4, G4R4, R5, G5R5 in which G3-G5 = defined as above and R3-R5 = independently alkyl substituted with 1-2 substituents selected from CO2H and derivs., CONH2 and derivs., SO3H, SO2NH2 and derivs.; A = CR6R7, CO, C:N-OH, C:N-OR8; R6, R7 = independently H, OH, OR8, alkyl; R8 = independently alkyl substituted with an aryl or cycloalkyl group; D = CH2, CHY; Y = O- or S-heterocycle; with the exclusion of compds. I in which A = CH2 and at least 3 of X1-X5 = H; and their stereoisomers, racemates, geometrical isomers, tautomers, salts, hydrates, solvates, solid forms and their mixts.] were prepared as PPAR activators, especially agonists, for treating dyslipidemia, diabetes type II and related diseases. Thus, reduction of 2-[2,6-dimethyl-4-[3-[4-(methylthio)phenyl]-3-oxoprop-1-enyl]phenoxy]-2methylpropanoic acid with triethylsilane in TFA at room temperature gave acid II (m.p. = 109-110°). Selected I were hPPAR α , hPPAR γ , and/or hPPA δ activators in an induced luciferase activity via hPPAR α /Gal4, hPPAR γ /Gal4, and hPPAR δ /Gal4 transactivation assay. I displayed hypolipemic properties by lowering the plasmatic cholesterol and triglycerides rates. I are useful for treating diabetes type II, dyslipidemia, pathologies associated with metabolic syndrome, cardiovascular diseases, etc.

IT 1000335-14-7, 2-[3-[4-(Methylthio)phenyl]-3-oxoprop-1-enyl]phenoxy]-2-methylpropanoic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of 1,3-diphenylpropane derivs. as PPAR activators for treating diseases especially dyslipidemia)

RN 1000335-14-7 HCAPLUS

CN Propanoic acid, 2-methyl-2-[3-[3-[4-(methylthio)phenyl]-3-oxo-1-propen-1-yl]phenoxy]- (CA INDEX NAME)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN 2007:1470010 HCAPLUS Full-text ACCESSION NUMBER:

DOCUMENT NUMBER: 148:100384

TITLE: Preparation of 1,3-diphenylpropane derivatives,

> particularly 2-[4-(3-oxo-3-phenylpropyl)phenoxy]-2methylpropanoic acids and related derivatives, as PPAR agonists for treating diseases especially dyslipidemia

Delhomel, Jean-Francois; Hanf, Remy; Caumont-Bertrand, INVENTOR(S):

Karine

PATENT ASSIGNEE(S): Genfit, Fr.

SOURCE: PCT Int. Appl., 97pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent French LANGUAGE:

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

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										WO 2	007-	EP56:	225	Ī	W 2	0070	621
OTHER S	OURCE	(S):			MAR	PAT	148:	1003	34								

GΙ

Title compds. I [X1 = halo, R1, G1R1; X2 = halo, R2, G2R2; X3 = R3, G3R3; X4 = halo, AB R4, G4R4; X5 = R5, G5R5; R1 = haloalkyl; R2 = H, alkyl; R3-R5 = independently H, (un) substituted alkyl; G1-G5 = independently O, S; with at least one of X3-X5 = R3, G3R3, R4, G4R4, R5, G5R5 in which G3-G5 = defined as above and R3-R5 = independently alkyl substituted with 1-2 substituents selected from CO2H and derivs., CONH2 and derivs., SO3H, SO2NH2 and derivs.; A = CR6R7, CO, C:N-OH, C:N-OR8; R6 = H, alkyl, OR8; R7 = alkyl, OH, OR8; R8 = independently alkyl substituted with an aryl or cycloalkyl group; D = CH2, CHY; Y = O- or S-heterocycle; and their stereoisomers, racemates, geometrical isomers, tautomers, salts, hydrates, solvates, solid forms and their mixts.] were prepared as PPAR activators, especially agonists, for treating dyslipidemia, diabetes type II and related diseases. Thus, reduction of 2-[2,6-dimethyl-4-[3-[4-(trifluoromethylthio)phenyl]-3-oxoprop-1-enyl]phenoxy]-2methylpropanoic acid with triethylsilane in DCM in the presence of TFA at room temperature gave the acid II (m.p. = $83-85^{\circ}$). Selected I were hPPAR α , hPPAR γ , and/or hPPA δ activators in an induced luciferase activity via hPPAR α /Gal4, hPPAR γ /Gal4, and hPPAR δ /Gal4 transactivation assay. I displayed hypolipemic properties by lowering the plasmatic cholesterol and triglycerides rates. I are useful for treating diabetes type II, dyslipidemia, pathologies associated with metabolic syndrome, cardiovascular diseases, etc.

IT 1000336-61-7, 2-[[4-[3-(4-Chloro-2-hydroxyphenyl)-3-oxoprop-1-enyl]phenyl]thio]-2-methylpropanoic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of 1,3-diphenylpropane derivs. as PPAR activators for treating diseases especially dyslipidemia) $\,$

RN 1000336-61-7 HCAPLUS

CN Propanoic acid, 2-[[4-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenyl]thio]-2-methyl- (CA INDEX NAME)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2006:1174214 HCAPLUS Full-text

DOCUMENT NUMBER: 145:483778

TITLE: Chalcones as farnesoid x receptor activators and

health foods

INVENTOR(S): Nozawa, Hajime

PATENT ASSIGNEE(S): Kirin Brewery Co., Ltd., Japan SOURCE: Jpn. Kokai Tokkyo Koho, 21pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2006306800 PRIORITY APPLN. INFO.:	A	20061109	JP 2005-132695 JP 2005-132695	20050428 20050428

OTHER SOURCE(S): MARPAT 145:483778

AB Chalcones, including xanthohumol from hop exts., are claimed as farnesoid x receptor (FXR) activators, adiponectin enhancers, and health foods for treatment of FXR-related diseases, including lipid metabolic diseases, diabetes, obesity, choledocholithiasis, fatty liver, hyperlipidemia, atherosclerosis, and other cardiovascular diseases, etc. The pharmacol. of xanthohumol were tested in animals.

IT 94-41-70, Chalcone, derivs.

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(chalcones as farnesoid x receptor activators and health foods)

RN 94-41-7 HCAPLUS

CN 2-Propen-1-one, 1,3-diphenyl- (CA INDEX NAME)

L12 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:19750 HCAPLUS Full-text

DOCUMENT NUMBER: 140:76896

TITLE: Composition based on substituted

1,3-diphenylprop-en-1-one derivatives, preparation and

use as PPARlpha agonists, antioxidants as well as

antiinflammatory agents

INVENTOR(S): Najib, Jamila; Caumont Bertrand, Karine

PATENT ASSIGNEE(S): Genfit S.A., Fr. SOURCE: Fr. Demande, 66 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
FR 2841784	A1	20040109	FR 2002-8570	20020708
FR 2841784	В1	20070302		
CA 2490993	A1	20040115	CA 2003-2490993	20030708
WO 2004005243	A2	20040115	WO 2003-FR2128	20030708
WO 2004005243	A3	20040422		

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            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
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                                                            A 20020708
PRIORITY APPLN. INFO.:
                                          FR 2002-8570
                                          WO 2003-FR2128 W 20030708
OTHER SOURCE(S):
                   MARPAT 140:76896
GΙ
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* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Title compds. I [wherein X1 = halo, R1, G1R1; X2 = H, thionitroso, OH, AB alkylcarbonyloxy, alkyloxy, SH, alkylthio, alkylcarbonylthio or X2 = O or S that forms a 2-phenyl-4H-1-benzopyran-4-one with the carbon-3 of the propene chain; X3 =R3, G3R3; X4 = halo, thionitroso, R4, G4R4; X5 = R5, G5R5; X6 = O, NH and derivs.; R1, R3, R4, R5 = independently H, (un)substituted alkyl; G1, G3, G4, G5 = independently O or S; with at least one of X1, X3, X4, or X5 of formula GR and one of the R1, R3, R4, or R5 is a substituted radical, and that radical form a cycle, or is associated with a group G; their optical and geometrical isomers, racemates, tautomers, salts, hydrates and mixts.; with the exclusion of certain compds.] were prepared as peroxisome proliferator-activated receptors- α (PPAR α) agonists and as antioxidants for treating cerebral ischemia and related diseases. For example, II was prepared by mixed-Aldol condensation of ketone III with 4-hydroxy-3,5ditertbutylbenzaldehyde in the presence of ethanol/HCl. In an antioxidant test, selected I (10-3 M) diminished the formation of oxidation product of LDL by AAPH by Selected I were PPARlpha agonists, showing induced luciferase activity via PPAR α /Gal4 transactivation with a factor of induction ranging from 10 to 60, 5-50 and 3-35 at 100 μM , 30 μM , and 10 μM resp. I and their compns. are useful for treating cardiovascular diseases, syndrome X, restenosis, diabetes, obesity, hypertension, inflammatory diseases, cancers or neoplasms (benign or malignant tumors), neurodegenerative diseases, dermatol. and the disorders related to the oxydative stress, for preventing and treating aging, and in particular cutaneous aging.

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IT 639864-16-7P 639864-17-8P 639864-18-9P 639864-19-0P 639864-20-3P 639864-21-4P 639864-22-5P 639864-23-6P 639864-30-5P
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639864-31-6P 639864-38-3P 639864-39-4P 639864-40-7P 639864-41-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(PPAR α agonist; preparation of diphenylpropenones as PPAR agonists for treating ischemia)

RN 639864-16-7 HCAPLUS

CN Propanoic acid, 2-[3-(1,1-dimethylethyl)-2-hydroxy-5-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl- (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Ho}_2\text{C} \xrightarrow{\text{C}} \text{O} \\ \text{Me} \\ \text{C} \xrightarrow{\text{CH}} \text{CH} \xrightarrow{\text{CH}} \text{Bu-t} \end{array}$$

RN 639864-17-8 HCAPLUS

CN Propanoic acid, 2-[3-(1,1-dimethylethyl)-2-hydroxy-5-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-18-9 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-3-(1,1-dimethylethyl)-2-hydroxyphenoxy]-2-methyl- (CA INDEX NAME)

RN 639864-19-0 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-3-(1,1-dimethylethyl)-2-hydroxyphenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

$$\begin{array}{c|c} \text{OH} & \text{O} \\ \hline \\ \text{C} & \text{CH} \\ \hline \\ \text{i-Pro-C} & \text{OH} \\ \hline \\ \text{O} & \text{Me} \\ \end{array}$$

RN 639864-20-3 HCAPLUS

CN Benzeneacetic acid, $3-(1,1-dimethylethyl)-2-hydroxy-5-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-<math>\alpha$, α -dimethyl- (CA INDEX NAME)

RN 639864-21-4 HCAPLUS

CN Benzeneacetic acid, $3-(1,1-\text{dimethylethyl})-2-\text{hydroxy}-5-[3-(2-\text{hydroxyphenyl})-3-\text{oxo}-1-\text{propen}-1-\text{yl}]-\alpha,\alpha-\text{dimethyl}-$, 1-methylethyl ester (CA INDEX NAME)

RN 639864-22-5 HCAPLUS

CN Benzeneacetic acid, $5-[3-(4-\text{chloro}-2-\text{hydroxyphenyl})-3-\text{oxo}-1-\text{propen}-1-\text{yl}]-3-(1,1-\text{dimethylethyl})-2-\text{hydroxy}-\alpha,\alpha-\text{dimethyl}-$ (CA INDEX NAME)

$$C1$$

OH

OH

CH

CH

CH

CH

CH

OH

Me

CCO2H

Me

RN 639864-23-6 HCAPLUS

CN Benzeneacetic acid, $5-[3-(4-\text{chloro}-2-\text{hydroxyphenyl})-3-\text{oxo}-1-\text{propen}-1-\text{yl}]-3-(1,1-\text{dimethylethyl})-2-\text{hydroxy}-\alpha,\alpha-\text{dimethyl}-$, 1-methylethyl ester (CA INDEX NAME)

RN 639864-30-5 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-2,3-dihydroxyphenoxy]-2-methyl- (CA INDEX NAME)

RN 639864-31-6 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-2,3-dihydroxyphenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-38-3 HCAPLUS

CN Propanoic acid, 2-[3-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl- (CA INDEX NAME)

RN 639864-39-4 HCAPLUS

CN Propanoic acid, 2-[3-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-40-7 HCAPLUS

CN Propanoic acid, 2-[[4-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenyl]thio]-2-methyl- (CA INDEX NAME)

RN 639864-41-8 HCAPLUS

CN Propanoic acid, 2-[[4-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenyl]thio]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

REFERENCE COUNT:

THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> => FILE HCAPLUS

FILE 'HCAPLUS' ENTERED AT 12:04:33 ON 30 JUN 2009
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FILE COVERS 1907 - 30 Jun 2009 VOL 151 ISS 1

FILE LAST UPDATED: 29 Jun 2009 (20090629/ED)

REVISED CLASS FIELDS (/NCL) LAST RELOADED: Apr 2009

USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Apr 2009

HCAplus now includes complete International Patent Classification (IPC) reclassification data for the second quarter of 2009.

CAS Information Use Policies apply and are available at:

http://www.cas.org/legal/infopolicy.html

This file contains CAS Registry Numbers for easy and accurate substance identification.
'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

=> D STAT QUE L18

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L18 ANSWER 1 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2006:491764 HCAPLUS Full-text

DOCUMENT NUMBER: 145:1047

TITLE: Methods and compositions using sirtuin modulators for

treating or preventing obesity and insulin resistance

disorders

INVENTOR(S): Sinclair, David A.; Alexander-Bridges, Maria

PATENT ASSIGNEE(S): President and Fellows of Harvard College, USA; The

General Hospital Corporation

SOURCE: U.S. Pat. Appl. Publ., 154 pp., Cont.-in-part of U.S.

Ser. No. 27,779.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PAT	FENT	NO.			KIN	D	DATE			APPL	ICAT	ION	NO.		D	ATE		
US	2006	0111	 435		A1	_	 2006	 0525		 US 2	2005-	 1740	00		2	0050	701	<
US	2005	0171	027		A1		2005	0804		US 2	004-	2777	9		2	0041	229	<
AU	2006	2661	25		A1		2007	0111		AU 2	006-	2661	25		2	0060	628	
CA	2613	636			A1		2007	0111		CA 2	006-	2613	636		2	0060	628	
WO	2007	0054	53		A2		2007	0111		WO 2	006-	US25	138		2	0060	628	
WO	2007	0054	53		АЗ		2007	0614										
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											004-					0040		
											2004-		-					
											2005-					0050		
										WO 2	006-	US25	T38		W 2	0060	628	

- AB The invention provides methods and compns. for modulating the activity or level of a sirtuin, thereby treating or preventing obesity or an insulin resistance disorder, e.g. diabetes, in a subject. Exemplary methods comprise contacting a cell with a sirtuin activating compound or an inhibitory compound to thereby increase or decrease fat accumulation, resp.
- IT 94-41-7, Chalcone 961-29-5, Isoliquiritigenin

13745-20-5, 4,2',4'-Trihydroxychalcone

RL: PAC (Pharmacological activity); BIOL (Biological study) (sirtuin modulators for treatment or prevention of obesity and insulin resistance disorders)

RN 94-41-7 HCAPLUS

CN 2-Propen-1-one, 1,3-diphenyl- (CA INDEX NAME)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

RN 13745-20-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)- (CA INDEX NAME)

IT 94-41-7D, Chalcone, derivs. 487-52-5, Butein

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

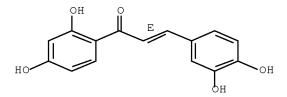
RN 94-41-7 HCAPLUS

CN 2-Propen-1-one, 1,3-diphenyl- (CA INDEX NAME)

RN 487-52-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(3,4-dihydroxyphenyl)-, (2E)-(CA INDEX NAME)

Double bond geometry as shown.



L18 ANSWER 2 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:638724 HCAPLUS Full-text

DOCUMENT NUMBER: 143:126796

TITLE: Compositions using sirtuin modulators for treating or

preventing obesity and insulin resistance disorders

INVENTOR(S): Sinclair, David A.; Alexander-Bridges, Maria

PATENT ASSIGNEE(S): President and Fellows of Harvard College, USA; The

General Hospital Corporation

SOURCE: PCT Int. Appl., 110 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

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		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KΖ,	LC,	
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EP	1708	689			A2		2006	1011		EP 2	004-	8158	42		2	0041	229	<
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CN	1012	4779.	3		А		2008	0820		CN 2	004-	8003	9345		2	0060	629	<
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										WO 2	004-	US43	847		W 2	0041	229	
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AB Methods and compns. are provided for modulating the activity or level of a sirtuin, thereby treating or preventing obesity or an insulin resistance disorder, e.g. diabetes, in a subject. Exemplary methods comprise contacting a cell with a sirtuin activating compound or an inhibitory compound to thereby increase or decrease fat accumulation, resp.

IT 487-52-5, Butein

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

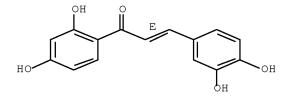
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(sirtuin modulators for treatment or prevention of obesity and insulin resistance disorders)

RN 487-52-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(3,4-dihydroxyphenyl)-, (2E)-(CA INDEX NAME)

Double bond geometry as shown.



L18 ANSWER 3 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2005:409480 HCAPLUS Full-text

DOCUMENT NUMBER: 142:463610

TITLE: Preparation of pyridines as inhibitors of dipeptidyl

peptidase IV useful for the prophylaxis or treatment

of diabetes

INVENTOR(S): Oi, Satoru; Maezaki, Hironobu; Suzuki, Nobuhiro PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan

SOURCE: PCT Int. Appl., 431 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA:	ΓΕΝΤ	NO.			KIN	D	DATE			APPL	ICAT	ION :	NO.		D	ATE		
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EP	1678	138			A1		2006	0712		EP 2	004-	7933	77		2	0041	029	<
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙΤ,	LI,	LU,	NL,	SE,	MC,	PT,	
		ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	PL,	SK,	HR
CN	1886	376			Α		2006	1227	1	CN 2	004-	8003	4965		2	0041	029	<
BR	2004	0159	60		Α		2007	0116		BR 2	004-	1596	0		2	0041	029	<
ZA	2006	0031	53		Α		2007	0829		ZA 2	006-	3153			2	0041	029	<
RU				C2		2009	0427		RU 2	006-	1188	06		2	0041	029	<	
MX	IX 2006003979				Α		2006	0705		MX 2	006-	3979			2	0600	407	<
US	2007	0037	807		A1		2007	0215		US 2	006-	5775	61		2	0000	428	<

Ser	ial	#•	10	/520,	078
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2006064022	A	20060612	KR	2006-708423		20060429	<
858259	B1	20080911					
2006KN01220	A	20070427	IN	2006-KN1220		20060510	<
2006002516	A	20060725	ИО	2006-2516		20060531	<
2008067013	A	20080717	KR	2008-715446		20080625	<
Y APPLN. INFO.:			JP	2003-373776	Α	20031031	<
			JΡ	2004-30491	Α	20040206	
			JΡ	2004-165977	Α	20040603	
			WO	2004-JP16457	W	20041029	
			KR	2006-708423	АЗ	20060429	
	858259 2006KN01220 2006002516 2008067013	858259 B1 2006KN01220 A 2006002516 A 2008067013 A	858259 B1 20080911 2006KN01220 A 20070427 2006002516 A 20060725 2008067013 A 20080717	858259 B1 20080911 2006KN01220 A 20070427 IN 2006002516 A 20060725 NO 2008067013 A 20080717 KR Y APPLN. INFO.: JP JP JP	858259 B1 20080911 2006KN01220 A 20070427 IN 2006-KN1220 2006002516 A 20060725 NO 2006-2516 2008067013 A 20080717 KR 2008-715446	858259 B1 20080911 2006KN01220 A 20070427 IN 2006-KN1220 2006002516 A 20060725 NO 2006-2516 2008067013 A 20080717 KR 2008-715446 Y APPLN. INFO.: JP 2003-373776 A JP 2004-30491 A JP 2004-165977 A WO 2004-JP16457 W	858259 B1 20080911 2006KN01220 A 20070427 IN 2006-KN1220 20060510 2006002516 A 20060725 NO 2006-2516 20060531 2008067013 A 20080717 KR 2008-715446 20080625 Y APPLN. INFO.: JP 2004-30491 A 20040206 JP 2004-165977 A 20040603 WO 2004-JP16457 W 20041029

OTHER SOURCE(S): CASREACT 142:463610; MARPAT 142:463610

GΙ

AB Title compds. I [wherein R1, R2 = independently (un)substituted hydrocarbyl, hydroxy; R3 = (un)substituted aryl; R4 = NH2 and derivs.; L = divalent hydrocarbon chain; Q = a bond or a divalent hydrocarbon chain; X = H, CN, NO2, acyl, OH and derivs., SH and derivs., NH2 and derivs., (un)substituted cyclyl; provided that when X = -C(:0)OEt, then Q = divalent hydrocarbon chain and that certain compds. are absent; and their salts, prodrugs] were prepared as dipeptidyl peptidase IV inhibitors. For example, Boc-protection of Me 5-(aminomethyl)-6-isobutyl-2-methyl-4-(4- methylphenyl)nicotinate (preparation given), saponification, coupling of the acid with isobutylamine/deprotection gave II•2TFA. I show a superior dipeptidyl peptidase IV inhibitory activity, and are useful as agents for the prophylaxis or treatment of diabetes and related diseases.

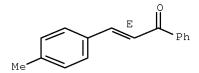
IT 22252-14-8P, (2E)-3-(4-Methylphenyl)-1-phenylprop-2-en-1-one
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)

(intermediate; preparation of pyridines as inhibitors of dipeptidyl peptidase IV useful for prophylaxis or treatment of diabetes)

RN 22252-14-8 HCAPLUS

CN 2-Propen-1-one, 3-(4-methylphenyl)-1-phenyl-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 4 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:993141 HCAPLUS Full-text

DOCUMENT NUMBER: 141:388723

TITLE: Flavonoid glycosides with enzymic modification for

prevention and treatment of type-II diabetes

INVENTOR(S): Tamura, Wataru; Matsuyama, Kayo; Kagami, Yoshiaki

PATENT ASSIGNEE(S): Ezaki Glico Co., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 6 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 2004323469	A	20041118	JP 2003-123434	20030428 <
PRIORITY APPLN. INFO.:			JP 2003-123434	20030428 <

AB Flavonoid glycosides, including flavane, flavanone, flavanol, flavone, isoflavone, and chalcone, with enzymic modification on their sugar chain are claimed as drugs and health foods for prevention and treatment of type-II diabetes.

IT 94-41-7, Chalcone

RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(flavonoid glycosides with enzymic modification for prevention and treatment of type-II diabetes)

RN 94-41-7 HCAPLUS

CN 2-Propen-1-one, 1,3-diphenyl- (CA INDEX NAME)

L18 ANSWER 5 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:902361 HCAPLUS $\underline{\text{Full-text}}$

DOCUMENT NUMBER: 141:395802

TITLE: Preparation of substituted phenylalkanoic acids,

including amino acid derivatives

INVENTOR(S): Van Zandt, Michael C.; Fang, Haiquan; Hu, Shaojing;

Whitehouse, Darren

PATENT ASSIGNEE(S): The Institutes for Pharmaceutical Discovery, LLC, USA

SOURCE: PCT Int. Appl., 131 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA.						KIND DATE		APPLICATION NO.						DATE			
	2004 2004				A2		2004	1028	,	WO 2	004-	US11	650		2	0040	414 <
	W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	AZ,	BA,	BB,	ВG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	KΖ,	LC,
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MΖ,	NA,	NI,
		NO,	NΖ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
		ΤJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW
	RW:	BW,	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	AZ,
		BY,	KG,	KΖ,	MD,	RU,	ΤJ,	TM,	ΑT,	BE,	ВG,	CH,	CY,	CZ,	DE,	DK,	EE,
		ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,
		SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,
		TD,	TG														
AU	2004	2311	06		A1		2004	1028		AU 2	004-	2311	06		2	0040	414 <
CA	2522	080			A1		2004	1028	1	CA 2	004-	2522	080		2	0040	414 <
US	2004	0248	937		A1		2004	1209		US 2	004-	8240	57		2	0040	414 <
EP	1633				A2 20060315				EP 2004-750170						20040414 <		
EP	1633	354			В1		2008	0123									
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙΤ,	LI,	LU,	NL,	SE,	MC,	PT,
		ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	PL,	SK, HR
BR	2004																414 <
	1794																414 <
JP	2006	5242	48		Τ		2006	1026	1	JP 2	006-	5100	73		2	0040	414 <
AT	3845	26			Τ		2008	0215		AT 2	004-	7501	70		2	0040	414 <
NO	2005	0047	69		Α		2006	0103		NO 2	005-	4769			2	0051	017 <
IN	2005	KN02	090		Α		2006	1117		IN 2	005-	KN20	90		2	0051	024 <
ZA	2005	0091	23		А		2007	0425		ZA 2	005-	9123			2	0051	111 <
PRIORIT	Y APP	LN.	INFO	.:						US 2	003-	4631	02P		P 2	0030	414 <
									,	WO 2	004-	US11	650	,	W 2	0040	414
OTHER SO	OURCE	(S):			MAR	PAT	141:	3958	02								

$$\begin{array}{c} Z \\ QL3 \end{array}$$

The invention relates to compds. I [n is 0-3; R1 is H, alkyl, phenylalkyl or alkenyl; R2 is Ph, phenylalkyl, alkyl, carbamoylalkyl, alkylsulfonylalkyl, heterocycloalkyl, etc.; R3 is H or CO2R1; R20-R23 are independently H, arylalkoxy, arylalkyl, halo, alkyl, OH, alkoxy, NO2, NH2, alkylamino, etc.; L is SO2NH, sulfonyl(alkylimino), NHSO2, O, CONH, carbonyl(alkylimino), SO2, carbonylalkylene, alkylenecarbonyl, NH or alkylimino (the alkyl group are optionally substituted with Ph or substituted phenyl); L2 is a bond, CONR9, NR9CO, alkylene-CONR9, NR9, etc. (R9 is H or alkyl optionally substituted with CO2H, arylsulfonyl or arylalkyl); ring A is (un)substituted Ph, naphthyl, thiazolyl, pyrazolyl, furanyl, dihydropyrazolyl,

benzofuranyl, dibenzofuranyl, pyrimidyl, pyridyl, quinolinyl, naphthyl, quinazolinyl, benzo[b]thiophene, imidazolyl, isothiazolyl, pyrrolyl, oxazolyl or triazolyl; Q is H, aryl, arylcarbonylaryl, alkyl, halo, etc.; L3 is a bond, alkyleneoxy, oxyalkylene, alkylene, alkenylene or CO; Z is absent, H, aroylamino, (un)substituted Ph or cycloalkylcycloalkanoyl(alkyl)amino] and their pharmaceutically-acceptable salts, which are useful in the treatment of metabolic disorders related to insulin resistance or hyperglycemia. These compds. include inhibitors of protein tyrosine phosphatase (PTP-1B) that are useful in the treatment of diabetes and other PTP-1B mediated diseases such as cancer and neurodegenerative diseases. Thus, 2-[4-[4-(4-chlorophenyl)-5-(4-ethylphenyl)thiazol-2-ylcarbamoyl]benzenesulfonylamino]-3-phenylpropionic acid was prepared by cyclocondensation of 4-ClC6H4COCH2C6H4Et-4 (preparation given) with thiourea, acylation with 4-ClSO2C6H4CO2H, and coupling with phenylalanine tert-Bu ester hydrochloride. The product was shown to increase the glucose infusion rate in rats at 30 mg/kg.

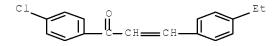
IT 782483-60-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of substituted phenylalkanoic acids, including amino acid derivs., for treatment of diabetes)

RN 782483-60-7 HCAPLUS

CN 2-Propen-1-one, 1-(4-chlorophenyl)-3-(4-ethylphenyl)- (CA INDEX NAME)



REFERENCE COUNT: 16 THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 6 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2003:896295 HCAPLUS Full-text

DOCUMENT NUMBER: 140:192745

TITLE: A licorice ethanolic extract with peroxisome

proliferator-activated receptor- γ ligand-binding activity affects diabetes in KK-Ay mice, abdominal

obesity in diet-induced obese C57BL mice and hypertension in spontaneously hypertensive rats Mae, Tatsumasa; Kishida, Hideyuki; Nishiyama, Tozo;

AUTHOR(S):

Mae, Tatsumasa; Kishida, Hideyuki; Nishiyama, Tozo;
Tsukagawa, Misuzu; Konishi, Eisaku; Kuroda, Minpei;
Mimaki, Yoshihiro; Sashida, Yutaka; Takahashi, Kazuma;

Kawada, Teruo; Nakagawa, Kaku; Kitahara, Mikio

CORPORATE SOURCE: Functional Foods Development Division, Life Science RD

Center, Kaneka Corporation, Hyogo, 676-8688, Japan

SOURCE: Journal of Nutrition (2003), 133(11),

3369-3377

CODEN: JONUAI; ISSN: 0022-3166

PUBLISHER: American Society for Nutritional Sciences

DOCUMENT TYPE: Journal LANGUAGE: English

AB The metabolic syndrome, including type 2 diabetes, insulin resistance, obesity/abdominal obesity, hypertension and dyslipidemia, is a major public health problem. Peroxisome proliferator-activated receptor-γ (PPAR-γ) ligands such as thiazolidinediones are effective against this syndrome. In this study, we showed that nonaq. fractions of licorice (Glycyrrhiza uralensis Fisher) extracted with ethanol, Et acetate and acetone, but not an aqueous extract, had PPAR-γ ligand-

binding activity with a GAL4-PPAR- γ chimera assay. Some prenylflavonoids including glycycoumarin, glycyrin, dehydroglyasperin C and dehydroglyasperin D, a newly found compound, were identified as active compds. with PPAR- γ ligand-binding activity in the nonaq. fraction of licorice. A licorice ethanolic extract contained these four active compds. at a total concentration of 16.7 g/100 g extract Feeding the licorice ethanolic extract at 0.1-0.3 g/100 g diet [.apprx.100 to 300 mg/(kg body·d)] for 4 wk decreased (P < 0.05) blood glucose level in younger (6 wk old) and older (13 wk old) diabetic KK-Ay mice and reduced (P < 0.05) wts. of intra-abdominal adipose tissues in high fat diet-induced obese C57BL mice. An increase in blood pressure in spontaneously hypertensive rats was suppressed (P < 0.01) by 3 wk of oral administration of the licorice ethanolic extract at 300 mg/(kg body·d). These findings indicate that licorice ethanolic extract is effective in preventing and ameliorating diabetes, ameliorating abdominal obesity and preventing hypertension, and suggest that licorice ethanolic extract would be effective in preventing and/or ameliorating the metabolic syndrome.

IT 961-29-5, Isoliquiritigenin

RL: NPO (Natural product occurrence); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses)

(licorice ethanolic extract with PPAR- γ ligand-binding activity affects diabetes in KK-Ay mice, abdominal obesity in diet-induced obese C57BL mice and hypertension in spontaneously hypertensive rats)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: 49 THERE ARE 49 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 7 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2002:487337 HCAPLUS Full-text

DOCUMENT NUMBER: 137:68154

TITLE: Method and composition for the treatment of diabetic

neuropathy

INVENTOR(S):
Rosenbloom, Richard A.

PATENT ASSIGNEE(S): The Quigley Corporation, USA

SOURCE: PCT Int. Appl., 31 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.		KIN	D	DATE			APPL	ICAT	ION I	. O <i>l</i>		D.	ATE	
			_									_		
WO 200204957	75	A2		2002	0627	•	WO 2	001-	US49.	297		2	0011	219 <
WO 200204957	15	А3		2003	0724									
W: AE,	AG, AL	, AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
CO,	CR, CU	, CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,

Serial#: 10/520,078 GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG US 20020115618 20020822 US 2000-740811 20001221 <--Α1 20030429 US 6555573 В2 US 20020165207 Α1 20021107 US 2001-847121 20010502 <--CA 2431079 Α1 20020627 CA 2001-2431079 20011219 <--AU 2002031095 20020701 AU 2002-31095 20011219 <--Α EP 1351679 20031015 EP 2001-991367 Α2 20011219 <--AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR Τ 20040603 JP 2002-550919 20011219 <--JP 2004516257 NZ 2001-526041 20011219 <--NZ 526041 Α 20050128 AU 2002231095 В2 20051124 AU 2002-231095 20011219 <--CA 2470603 Α1 20030703 CA 2002-2470603 20021106 <--WO 2003053336 Α2 20030703 WO 2002-US35654 20021106 <--WO 2003053336 АЗ 20031127 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, W: CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20030709 AU 2002352501 Α1 AU 2002-352501 20021106 <--EP 1455778 A2 20040915 EP 2002-789474 20021106 <--AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK Т 20050623 JP 2003-554096 JP 2005518381 20021106 <--

20060630

20030724

A 20040602 A 20070302 A 20041203 A 20050829 A 20070511 A 20040927

Α

Α1

WO 2002-US35654 W 20021106 <--AB A composition for the treatment of diabetic neuropathy comprise a mixture of a compound that promotes synthesis of nerve growth factor, an aldose reductase inhibitor and an antioxidant, optionally formulated in a pharmaceutically acceptable carrier. This combination of active agents provides significant, effective relief of the symptoms of diabetic neuropathy, as well as at least partial recovery of lost neurol. function in some cases. In addition, the compns. of the present invention, when used in effective amts. to treat diabetic neuropathy, do not exhibit the severe side effects of many prior art compns. proposed for treatment of this ailment. An effective amount of the composition of the invention is administered over a period of time sufficient to provide the beneficial effects of relief from the symptoms of diabetic neuropathy, as well as at least some recovery of the damaged nerve tissues. For example, A topical composition including a mixture of an hydrophilic ointment

NZ 2002-533439

US 2003-369025

ZA 2003-4247

IN 2003-DN870

MX 2003-5672

ZA 2004-4614

IN 2004-DN1683

MX 2004-6039

US 2000-740811

US 2001-847121

WO 2001-US49297

20021106 <--

20030219 <--

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20030620 <--

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A 20001221 <--

A 20010502 <--

W

NZ 533439

US 20030138504

IN 2003DN00870

MX 2003005672

ZA 2004004614

IN 2004DN01683

MX 2004006039

PRIORITY APPLN. INFO.:

ZA 2003004247

base, sodium acid phosphate moisturizing agent, witch hazel extract, glycerin, apricot kernel oil and DL-panthenol, together with pharmaceutically acceptable carrier, and further including, as active agents, vitamins A and D3, ascorbyl palmitate, quercetin and vitamin E acetate, was prepared by cold compounding. The topical composition was applied twice daily in the morning and afternoon under the supervision of a physician, but patients were permitted to apply the composition up to six times daily, as needed for pain relief over a period of a few days. All patients treated experienced immediate pos. results that lasted up to a day or two after treatment was discontinued. The effects noted by the patients included the relief of burning pain, tingling, healing of damaged skin, and reversal of skin discoloration due to impaired circulation.

IT 961-29-5, Isoliquiritigenin

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(compns. containing nerve growth factor promoters, aldose reductase inhibitors and antioxidants for treatment of diabetic

neuropathy)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 8 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2001:401646 HCAPLUS Full-text

DOCUMENT NUMBER: 135:152641

TITLE: Synthesis of flavonoids and their effects on aldose

reductase and sorbitol accumulation in streptozotocin-induced diabetic rat tissues

AUTHOR(S): Lim, Soon Sung; Jung, Sang Hoon; Ji, Jun; Shin, Kuk

Hyun; Keum, Sam Rok

CORPORATE SOURCE: Natural Products Research Institute, Seoul National

University, Seoul, S. Korea

SOURCE: Journal of Pharmacy and Pharmacology (2001),

53(5), 653-668

CODEN: JPPMAB; ISSN: 0022-3573

PUBLISHER: Pharmaceutical Press

DOCUMENT TYPE: Journal LANGUAGE: English

OTHER SOURCE(S): CASREACT 135:152641

AB The purpose of this study was to develop new compds. with these dual-effects through synthesis of chalcone derivs. and by examining the structure-activity relationships on the inhibition of rat lens aldose reductase as well as on antioxidant effects. A series of 35 flavonoid derivs. were synthesized by Winget's condensation, oxidation, and reduction of appropriate acetophenones with appropriate benzaldehydes. The inhibitory activity of these derivs. on rat lens aldose reductase and their antioxidant effects, measured using Cu2+ chelation and radical scavenging activities on 1,1-diphenyl-picrylhydrazyl in-vitro, were evaluated. Their effect on sorbitol

accumulation in the red blood cells, lenses and sciatic nerves of streptozotocininduced diabetic rats was also estimated Among the new flavonoid derivs. synthesized, those with the 2',4'-dihydroxyl groups in the A ring such as 2,4,2',4'tetrahydroxychalcone, 2,2',4'-trihydroxychalcone, 2',4'-dihydroxy-2,4dimethylchalcone and 3,4,2',4'-tetrahydroxychalcone (I) were found to possess the highest rat lens aldose reductase inhibitory activity in-vitro, their IC50 values (concentration of inhibitors giving 50% inhibition of enzyme activity) being 1.6 + 10-7, 3.8 + 10-7, 4.0 + 10-7 and 4.6 + 10-7 M, resp. All of the chalcones tested except those with o-dihydroxy or hydroquinone moiety showed a weak free radical scavenging activity. In the in-vivo expts., however, compound I with o-dihydroxy moiety in the B ring showed the strongest inhibitory activity in the accumulation of sorbitol in the tissues. It also showed the strongest activity in transition metal chelation and free radical scavenging activity. Of the 4,2'-dihydroxyl and 2',4'dihydroxyl derivs. of flavonoid synthesized, including chalcone, flavone, flavanone, flavonol and dihydrochalcone, some chalcone derivs. synthesized were found to possess aldose reductase inhibition and antioxidant activities in-vitro as well as inhibition in the accumulation of sorbitol in the tissues in-vivo. 3,4,2',4'-Tetrahydroxychalcone (I, butein) was the most promising compound for the prevention or treatment of diabetic complications.

IT 487-52-5P, Butein 961-29-5P 25515-43-9P 34000-31-2P 34000-35-6P 318296-33-2P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(synthesis of flavonoids and effects on aldose reductase and sorbitol accumulation in streptozotocin-induced diabetic rat tissues)

RN 487-52-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(3,4-dihydroxyphenyl)-, (2E)-(CA INDEX NAME)

Double bond geometry as shown.

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

RN 25515-43-9 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-phenyl-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

RN 34000-31-2 HCAPLUS

CN 2-Propen-1-one, 1-(2-hydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

RN 34000-35-6 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-methylphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

RN 318296-33-2 HCAPLUS

CN 2-Propen-1-one, 3-(4-bromophenyl)-1-(2,4-dihydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: 41 THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 9 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1997:563086 HCAPLUS Full-text

DOCUMENT NUMBER: 127:220466

ORIGINAL REFERENCE NO.: 127:42961a,42964a

TITLE: Preparation and formulation of phenylalkanediones and

analogs as therapeutic agents for diabetes

INVENTOR(S): Shinkai, Hisashi; Ozeki, Hidekazu; Furukawa, Noboru PATENT ASSIGNEE(S): Japan Tobacco Inc., Japan; Shinkai, Hisashi; Ozeki,

Hidekazu; Furukawa, Noboru

SOURCE: PCT Int. Appl., 220 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

								APPLICATION NO.										
								WO 1997-JP422										
	W:	AL,	AM,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	DE,	
		DK,	EE,	ES,	FI,	GB,	GE,	HU,	IL,	IS,	JP,	KΕ,	KG,	KR,	KΖ,	LC,	LK,	
		LR,	LS,	LT,	LU,	LV,	MD,	MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	
		RU,	SD,	SE,	SG,	SI,	SK,	ТJ,	TM,	TR,	TT,	UA,	UG,	US,	UZ,	VN,	YU	
	RW:	ΚE,	LS,	MW,	SD,	SZ,	UG,	ΑT,	BE,	CH,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	
		IE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	ML,	
		MR,	NE,	SN,	TD,	ΤG												
CA	2246	725			A1		1997	0821	1	CA 1	997-	2246	725		1	9970.	217	<
AU	9716	732			Α		1997	0902	-	AU 1	997-	1673	2		1	9970.	217	<
-	7193							0511										
JP	0928	6755			Α		1997	1104	1	JP 1	997-	4980	3		1	9970.	217	<
	3104							1030										
EP	8858	69			A1		1998	1223		EP 1	997-	9027	12		1	9970.	217	<
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙT,	LI,	LU,	NL,	SE,	MC,	PT,	
		IE,																
	1216							0512										
	9900							0628		HU 1	999-	715			1	9970.	217	<
	9900							1128										
	9707				А			0727								9970		
	2174				C2			0927		RU 1						9970.		
	1997							0304		IN 1			-			9970.		
	9803	· · · -			А		1998	1019		NO 1						9980		
RIORIT	Y APP	LN.	INFO	.:						JP 1			-			9960.		
									1	WO 1	997-	JP42	2		W 1	9970.	217	<

optionally substituted alkyl group having 1 to 5 carbon atoms, an optionally substituted aryl moiety, etc.; R2 represents a hydrogen atom, an optionally substituted alkyl group (having 1 to 5 carbon atoms), etc.; R represents a hydrogen atom; and R3 represents an optionally substituted alkyl group (having 1 to 5 carbon atoms), etc.] are prepared The title agents have excellent blood sugar lowering activity in the case of high blood sugar level, do not influence the blood sugar in the case of normal blood sugar level, i.e., do not cause any severe side effect, such as hypoglycemia, and are useful not only as therapeutic agents but also as prophylactics for chronic

complications of diabetes. 3-Benzoyl-1-cyclopentanone at 1 mg/kg gave 27.8% decrease

The title compds. R1COCRR2XCOR3 [wherein X represents O, etc.; R1 represents an

of blood sugar in rats dosed with glucose. IT 614-47-1, trans-Chalcone

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of phenylalkanediones and analogs as there

MARPAT 127:220466

(preparation of phenylalkanediones and analogs as therapeutic agents for diabetes)

RN 614-47-1 HCAPLUS

OTHER SOURCE(S):

AΒ

CN 2-Propen-1-one, 1,3-diphenyl-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L18 ANSWER 10 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1997:417489 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 127:130934

ORIGINAL REFERENCE NO.: 127:25125a,25128a

TITLE: Antioxidant constituents from licorice roots:

isolation, structure elucidation and antioxidative

capacity toward LDL oxidation

AUTHOR(S): Vaya, Jacob; Belinky, Paula A.; Aviram, Michael CORPORATE SOURCE: Migal, Galilee Technol. Cent., Kiryat Shmona, 10200,

Israel

SOURCE: Free Radical Biology & Medicine (1997),

23(2), 302-313

CODEN: FRBMEH; ISSN: 0891-5849

PUBLISHER: Elsevier
DOCUMENT TYPE: Journal
LANGUAGE: English

The present study analyzed the antioxidative properties of natural compds. from the root of the plant Glycyrrhiza glabra (licorice) toward LDL oxidation Seven constituents, with antioxidant capacity were isolated from Glycyrrhiza glabra. isolated compds. were identified as the isoflavans Hispaglabridin A, Hispaglabridin B, Glabridin, and 4'-0-Methylglabridin, the two chalcones, isoprenylchalcone derivative and Isoliquiritigenin, and the isoflavone, Formononetin. Among these compds., Glabridin constituted the major amount in the crude extract (11.6%, weight/weight) as detected by high-performance liquid chromatog. (HPLC) anal. The antioxidative capacities of the isolated compds. were tested against β -carotene destruction and LDL oxidation The isoflavans at a concentration of 50 μM inhibited β -carotene consumption, following 90 min of incubation at 50°, similar to the inhibitory effect of the whole licorice crude extract (at 16 mg/L). The chalcones exhibited moderate inhibition and the isoflavone was almost inactive whereas vitamin E (50 μ M) completely inhibited β -carotene consumption. The inhibitory effect of the constituents at a concentration of 30 μ M on 2,2'-azobis(2-amidinopropane) dihydrochloride (AAPH)-induced LDL oxidation was determined by measuring the amount of the thiobarbituric acid reactive substances (TBARS) and the amount of lipid peroxides. While the isoflavans and chalcones exhibited high inhibitory activity, Formononetin and vitamin E were not active. A dose-dependent inhibitory effect of Glabridin on the formation of cholesteryl linoleate hydroperoxide (CLOOH) in an AAPH-induced LDL oxidation system was also shown. Glabridin, at 5 or $40-60~\mu\mathrm{M}$ concentration, inhibited the CLOOH formation by 62% and 90%, resp. These results suggest that the isoflavans and chalcones are very potent antioxidants toward LDL oxidation with Glabridin being the most abundant and potent antioxidant. As LDL oxidation is a key event in the formation of the early atherosclerotic lesion, the use of these natural antioxidants may be proven beneficial to attenuate atherosclerosis.

IT 961-29-5P, Isoliquiritigenin

RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); PUR

(Purification or recovery); THU (Therapeutic use); BIOL (Biological

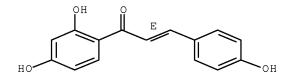
study); OCCU (Occurrence); PREP (Preparation); USES (Uses)

(antioxidant constituents from licorice roots and isolation and structure elucidation and antioxidative capacity toward LDL oxidation in relation to atheresclerosis inhibition)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.



L18 ANSWER 11 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1994:245079 HCAPLUS Full-text

DOCUMENT NUMBER: 120:245079

ORIGINAL REFERENCE NO.: 120:43453a,43456a

TITLE: Preparation of thiazolidine-2,4-dione derivatives as

antidiabetics

INVENTOR(S): Myaoka, Shozo; Sato, Hiroko; Takahashi, Keimei;

Ushijima, Hideto

PATENT ASSIGNEE(S): Terumo Corp, Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 7 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

GΙ

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 05310718	А	19931122	JP 1992-110460	19920428 <
PRIORITY APPLN. INFO.:			JP 1992-110460	19920428 <
OTHER SOURCE(S):	MARPAT	120:245079		

 $\begin{array}{c|c} \text{S} & \text{CH}_2 \\ \hline \\ \text{O} & \text{R}^2 \end{array}$

The title derivs. I (R1, R2 = H, OH, lower alkoxy, alkoxymethoxy) are prepared A mixture of 25.0 g 5-(3-acetylbenzyl)thiazolidine-2,4-dione (prepared from m-aminoacetophenone in 2 steps), 19.7 g 3-methoxy-4-methoxymethoxybenzaldehyde, and aqueous KOH in MeOH was treated at room temperature for 2.5 h to give 23.1 g 5-[3-(3-methoxy-4-methoxymethoxyphenyl)-2-propenoyl)benzyl]thiazolidine- 2,4-dione, which (6.20 g) was treated with tert-Bu bromacetate in the presence of K2CO3 in DMF

at room temperature for 1.5 h to give 3-tert-butoxycarbonylmethyl-5-[3-(3-(4-hydroxy-3-methoxyphenyl)-2- propencyl)benzyl]thiazolidine-2,4-dione (II). II (4.30 g) was stirred at room temperature in HCO2H for 2.5 h to give 2.70 g I (R1 = OMe, R2 = OH) (III). III inhibited aldose reductase with IC50 of 1.4 + 10-7.

IT 154066-97-4P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of, as antidiabetic)

RN 154066-97-4 HCAPLUS

CN 3-Thiazolidineacetic acid, 5-[[4-[3-(4-hydroxy-3-methoxyphenyl)-1-oxo-2-propen-1-yl]phenyl]methyl]-2,4-dioxo- (CA INDEX NAME)

PAGE 1-A

PAGE 2-A

L18 ANSWER 12 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1993:80926 HCAPLUS Full-text

DOCUMENT NUMBER: 118:80926

ORIGINAL REFERENCE NO.: 118:14241a,14244a

TITLE: Thiazolidine-2, 4-dione compounds, method for their

production, and medicines containing them for

treatment of diabetic complications

INVENTOR(S): Miyaoka, Shozo; Takahashi, Hiroaki; Ushijima, Hideto;

Sato, Hiroko

PATENT ASSIGNEE(S): Terumo Corp., Japan SOURCE: Eur. Pat. Appl., 11 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE		
				-			
EP 489663	A1	19920610	EP 1991-403312		19911206 <		
R: BE, CH, DE,	FR, GB	, IT, LI, NL	, SE				
JP 04210683	A	19920731	JP 1990-413602		19901206 <		
US 5225426	A	19930706	US 1991-802308		19911204 <		
PRIORITY APPLN. INFO.:			JP 1990-413602	Α	19901206 <		
OTHER SOURCE(S):	MARPAT	118:80926					
GI							

$$0 \underset{\text{HN}}{\overset{\circ}{\bigvee}} CH_2 \underset{\text{OH}}{\overset{\circ}{\bigvee}} OR^1$$

Title compds. I (R1 = H, Me) are prepared as antidiabetic drugs with a combined action, both inhibiting aldose reductase and depressing blood sugar. For example, condensation of 5-(4-acetylbenzyl)thiazolidine-2,4-dione (prepared in 2 steps) with 3,4-(MeO)(MeOCH2O)C6H3CHO in methanolic KOH, and deprotection of the product by HCl in aqueous THF-MeOH, gave I (R1 = Me; CH2 group in 4-position) (II). At 100 mg/kg/day orally for 4 days in diabetic rats, II gave 93.8% inhibition of sorbitol accumulation and 60.5% drop in blood sugar, whereas a comparative thiazoleacetic acid derivative gave 90.3% inhibition of sorbitol but only 7.2% blood sugar drop. Prepns. and biol. data for 3 I are described; these and 3 addnl. I are claimed.

IT 145704-63-8P 145704-64-9P 145704-67-2P

145704-68-3P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of, as antidiabetic)

RN 145704-63-8 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[4-[3-(3,4-dihydroxyphenyl)-1-oxo-2-propen-1-yl]phenyl]methyl]- (CA INDEX NAME)

PAGE 1-A

PAGE 2-A $_{\mathrm{OH}}$

RN 145704-64-9 HCAPLUS
CN 2,4-Thiazolidinedione, 5-[[4-[3-(4-hydroxy-3-methoxyphenyl)-1-oxo-2-propen-1-yl]phenyl]methyl]- (CA INDEX NAME)

PAGE 1-A

CH2

CH

CH

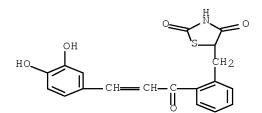
OMe

PAGE 2-A

 $\int_{\mathbb{H}}$

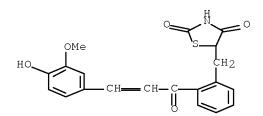
145704-67-2 HCAPLUS RN

 $2, 4- \\ Thiazolidine dione, 5-[[2-[3-(3,4-dihydroxyphenyl)-1-oxo-2-propen-1-dihydroxyphenyl])-1-oxo-2-propen-1-dihydroxyphenyl]$ CN yl]phenyl]methyl]- (CA INDEX NAME)



RM145704-68-3 HCAPLUS

CN 2,4-Thiazolidinedione, 5-[[2-[3-(4-hydroxy-3-methoxyphenyl)-1-oxo-2-propen-1-y1]phenyl]methyl]- (CA INDEX NAME)



L18 ANSWER 13 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 1992:51296 HCAPLUS Full-text

DOCUMENT NUMBER: 116:51296 ORIGINAL REFERENCE NO.: 116:8694h,8695a

TITLE: Effects of aldose reductase inhibitors on prostacyclin

(PGI2) synthesis by aortic rings from rats with

streptozotocin-induced diabetes

AUTHOR(S): Wakasugi, M.; Noguchi, T.; Inoue, M.; Tawata, M.;

Shindo, H.; Onaya, T.

CORPORATE SOURCE: Med. Sch., Univ. Yamanashi, Yamanashi, 409-38, Japan SOURCE:

Prostaglandins, Leukotrienes and Essential Fatty Acids

(1991), 44(4), 233-6

CODEN: PLEAEU; ISSN: 0952-3278

DOCUMENT TYPE: Journal LANGUAGE: English

AB The effects of aldose reductase inhibitors (ARIs) on the synthesis of PGI2 by aortic rings from diabetic rats were examined The ARIs studied were ONO-2235 and isoliquiritigenin, a new compound extracted from glycyrrhizae radix. The content of sorbitol in the sciatic nerve of diabetic rats induced by streptozotocin was increased as compared with that of controls. This increase was inhibited by the

administration of an ARI. On the other hand, there was a decrease in the synthesis of PGI2 by the diabetic rats compared with the control rats. The decrease in PGI2 synthesis was reversed by the administration of an ARI. Furthermore, the synthesis of PGI2 by the aortic rings was inversely correlated with the content of sorbitol in sciatic nerves. Those observations suggest that an ARI may have a beneficial effect on the vascular synthesis of PGI2 in diabetes mellitus.

IT 961-29-5, Isoliquiritigenin

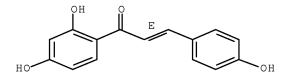
RL: BIOL (Biological study)

(aldose reductase inhibition by, prostacyclin formation in aorta response to, in diabetes mellitus)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.



L18 ANSWER 14 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1990:584545 HCAPLUS Full-text

DOCUMENT NUMBER: 113:184545

ORIGINAL REFERENCE NO.: 113:31051a,31054a

TITLE: Isoliquiritigenin: a new aldose reductase inhibitor

from Glycyrrhizae Radix

AUTHOR(S): Aida, Kaoru; Tawata, Masato; Shindo, Hideo; Onaya,

Toshimasa; Sasaki, Hiroshi; Yamaguchi, Takuji; Chin,

Masao; Mitsuhashi, Hiroshi

CORPORATE SOURCE: Med. Sch., Univ. Yamanashi, Yamanashi, 409-38, Japan

SOURCE: Planta Medica (1990), 56(3), 254-8

CODEN: PLMEAA; ISSN: 0032-0943

DOCUMENT TYPE: Journal LANGUAGE: English

GI

AB Traditionally in Japan, some kampo medicines (traditional oriental herbal prescriptions) have long been used for the treatment of diabetic neuropathy. Some aldose reductase inhibitors are included among these drugs. Thus, the components of Glycyrrhizae Radix, a constituent of some kampo medicines were studied and 6 compds. (GUs 9-17) were isolated. Among these, GU-17, identified as isoliquiritigenin (I), had the most potent aldose reductase inhibiting activity. I inhibited rat lens aldose reductase with an IC50 of 3.2 + 10-7 M, using DL-glyceraldehyde as a substrate. It inhibited sorbitol accumulation in human red blood cells in vitro,

with an IC50 of 2.0 + 10-6 M. I, when administered via an intragastric tube to diabetic rats, suppressed sorbitol accumulation in the red blood cells, the sciatic nerve, and the lens as effectively as ONO-2235. These results suggest that I may be effective in preventing diabetic complications.

IT 961-29-5, Isoliquiritigenin

RL: BIOL (Biological study)

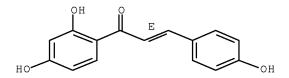
(of glycyrrhiza root, aldose reductase inhibition by, diabetes

in relation to)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.



L18 ANSWER 15 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1988:637045 HCAPLUS Full-text

DOCUMENT NUMBER: 109:237045

ORIGINAL REFERENCE NO.: 109:39113a,39116a

TITLE: Pharmaceuticals containing aldose reductase inhibitors

for treatment of diseases caused by diabetes

INVENTOR(S): Meya, Toshimasa; Tawada, Masato; Sasaki, Hiroshi;

Nishimura, Hiroaki

Ι

PATENT ASSIGNEE(S): Tsumura Juntendo, Inc., Japan SOURCE: Jpn. Kokai Tokkyo Koho, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 63104912	А	19880510	JP 1986-248389	19861021 <
JP 07055902	В	19950614		
PRIORITY APPLN. INFO.:			JP 1986-248389	19861021 <
OTHER SOURCE(S):	MARPAT	109:237045		
GT				

AΒ Pharmaceuticals contain aldose reductase inhibitors dihydroxyphenyl (phenyloxy) propenone derivs. (I; R = H, glucosyl, or apioglucosyl) for treatment of diseases derived from diabetes. I (R = glucosyl) was extracted from licorice, and purified by a series of column chromatog. I (R = glucosyl) 100 and anhydrous silicic acid 20 g were mixed, and 75 g corn starch was added, followed by 100 mL 10% hydroxypropyl cellulose-alc. mixture This mixture was made into granules.

ΙT 961-29-5

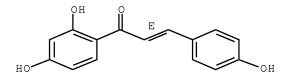
> RL: BIOL (Biological study) (pharmaceutical containing, for treatment of diseases related to

diabetes)

RN 961-29-5 HCAPLUS

CN 2-Propen-1-one, 1-(2,4-dihydroxyphenyl)-3-(4-hydroxyphenyl)-, (2E)- (CA)

Double bond geometry as shown.



L18 ANSWER 16 OF 16 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 1988:112221 HCAPLUS Full-text

DOCUMENT NUMBER: 108:112221

ORIGINAL REFERENCE NO.: 108:18373a, 18376a

TITLE: Preparation of (heterocyclylalkenyl) mevalonates as

> hypolipemics and antiatherosclerotic agents Wareing, James Richard; Damon, Robert Edson

PATENT ASSIGNEE(S): Sandoz A.-G., Switz.; Sandoz-Patent-G.m.b.H.; Sandoz-Erfindungen Verwaltungsgesellschaft m.b.H.

Eur. Pat. Appl., 41 pp. SOURCE:

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

INVENTOR(S):

PA:	TENT NO.			KINI	D DATE	APPLICATION NO.		DATE	
EP	221025			A1			_	19861021 <	-
1.10			CH,	•		GR, IT, LI, LU, NL, SE		10061001	
WO	8702662			A2	19870507	WO 1986-EP598		19861021 <	-
WO	8702662			А3	19871217				
	W: AU	, DK,	FI,	HU,	JP, KR				
AU	8665994			Α	19870519	AU 1986-65994		19861021 <	-
AU	598775			В2	19900705				
JP	6350115	3		Τ	19880428	JP 1986-505883		19861021 <	-
HU	48208			A2	19890529	HU 1986-5313		19861021 <	-
IL	80403			Α	19900917	IL 1986-80403		19861023 <	-
CA	1278794			С	19910108	CA 1986-521333		19861024 <	-
PL	154130			В1	19910731	PL 1986-262032		19861024 <	-
FΙ	8702299			Α	19870525	FI 1987-2299		19870525 <	-
DK	8703218			Α	19870624	DK 1987-3218		19870624 <	-
PRIORIT	Y APPLN.	INFO	.:			US 1985-791198	А	19851025 <	-

US 1986-816664 A 19860107 <--WO 1986-EP598 A 19861021 <--

OTHER SOURCE(S): MARPAT 108:112221

GI

The title compds. [I, II; R1,R2 = alkyl, cycloalkyl, (un)substituted Ph; R3 = R4, alkenyl; R4 = H, R1; X = (CH2)m, alkenylene; Y = NR4, O, S; Z = CHOHCH2CR5OHCH2CO2H; R5 = H, alkyl; m = 0-3] were prepared as hypolipemics and antiatherosclerotic agents (no data). PhCOCH2CH(CHMe2)COCO2Et (preparation given) and 4-FC6H4NH2 were refluxed 16 h in PhMe containing TiCl4 to give III (R = CO2Et) which was converted in 7 steps to (±)-erythro-III (R = CH:CHCHOHCH2CHOHCH2CO2Et).

IT 22966-07-0P

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation and condensation of, in preparation of hypolipemic and antiatherosclerotic agents)

RN 22966-07-0 HCAPLUS

CN 2-Propen-1-one, 3-(4-fluorophenyl)-1-phenyl-, (2E)- (CA INDEX NAME)

Double bond geometry as shown.

Serial#: 10/520,078 INVENTOR SEARCH

=> FILE HCAPLUS

FILE 'HCAPLUS' ENTERED AT 13:55:55 ON 30 JUN 2009
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PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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=> D STAT QUE L23	
L3 STR	
L4 (47768) SEA FILE=REGISTRY SSS	FUL L3
L5 STR	
L6 2679 SEA FILE=REGISTRY SUB-	=L4 SSS FUL L5
L7 9264 SEA FILE=HCAPLUS SPE=0	ON ABB=ON PLU=ON L6
L8 191239 SEA FILE=HCAPLUS SPE=C	ON ABB=ON PLU=ON (DIABETES/CT OR
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OR ?ATHEROSCLER?/BI	
L10 187726 SEA FILE=HCAPLUS SPE=C	ON ABB=ON PLU=ON OBESITY+NT/CT OR
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OR REDUCTION OR MANAGE	EMENT))/BI
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L12 5 SEA FILE=HCAPLUS SPE=C	ON ABB=ON PLU=ON L7 AND L11
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K?/AU OR CAUMONT K?/AU	U OR BERTRAND K?/AU
L21 5 SEA FILE=HCAPLUS SPE=C	ON ABB=ON PLU=ON L19 AND L20
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OR L20)	
L23 7 SEA FILE=HCAPLUS SPE=0	ON ABB=ON PLU=ON L21 OR L22

=> D L23 IBIB ABS HITSTR 1-7

L23 ANSWER 1 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2007:1470011 HCAPLUS Full-text

DOCUMENT NUMBER: 148:100385

TITLE: Preparation of 1,3-diphenylpropane derivatives,

particularly 2-[4-(3-oxo-3-phenylpropyl)phenoxy]-2methylpropanoic acids and related derivatives, as PPAR agonists for treating diseases especially dyslipidemia

INVENTOR(S):
Delhomel, Jean-Francois; Hanf, Remy;

Caumont-Bertrand, Karine

PATENT ASSIGNEE(S): Genfit, Fr.

SOURCE: PCT Int. Appl., 108pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: French FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

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         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA,
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PRIORITY APPLN. INFO.:
                                            FR 2006-5540
                                                                A 20060621
                                                               W 20070621
                                            WO 2007-EP56224
OTHER SOURCE(S):
                        MARPAT 148:100385
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Title compds. I [X1 = R1, G1R1; X2 = halo, R2, G2R2; X3 = R3, G3R3; X4 = halo, R4, G4R4; X5 = R5, G5R5; R1 = H, nonhalogenated alkyl; R2 = H, alkyl; R3-R5 = independently H, (un)substituted alkyl; G1-G5 = independently O, S; with at least one of X3-X5 = R3, G3R3, R4, G4R4, R5, G5R5 in which G3-G5 = defined as above and R3-R5 = independently alkyl substituted with 1-2 substituents selected from CO2H and derivs., CONH2 and derivs., SO3H, SO2NH2 and derivs.; A = CR6R7, CO, C:N-OH, C:N-OR8; R6, R7 = independently H, OH, OR8, alkyl; R8 = independently alkyl substituted with an aryl or cycloalkyl group; D = CH2, CHY; Y = O- or S-heterocycle; with the exclusion of compds. I in which A = CH2 and at least 3 of X1-X5 = H; and their stereoisomers, racemates, geometrical isomers, tautomers, salts, hydrates, solvates, solid forms and their mixts.] were prepared as PPAR activators, especially agonists, for treating dyslipidemia, diabetes type II and related diseases. Thus, reduction of 2-[2,6-dimethyl-4-[3-[4-(methylthio)phenyl]-3-oxoprop-1-enyl]phenoxy]-2-methylpropanoic acid with triethylsilane in TFA at room temperature gave acid II

GT

(m.p. = 109-110°). Selected I were hPPAR α , hPPAR γ , and/or hPPA δ activators in an induced luciferase activity via hPPAR α /Gal4, hPPAR γ /Gal4, and hPPAR δ /Gal4 transactivation assay. I displayed hypolipemic properties by lowering the plasmatic cholesterol and triglycerides rates. I are useful for treating diabetes type II, dyslipidemia, pathologies associated with metabolic syndrome, cardiovascular diseases, etc.

IT 1000335-14-7, 2-[3-[4-(Methylthio)phenyl]-3-oxoprop-1-

enyl]phenoxy]-2-methylpropanoic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of 1,3-diphenylpropane derivs. as PPAR activators for treating diseases especially dyslipidemia)

RN 1000335-14-7 HCAPLUS

CN Propanoic acid, 2-methyl-2-[3-[3-[4-(methylthio)phenyl]-3-oxo-1-propen-1-yl]phenoxy]- (CA INDEX NAME)

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 2 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2007:1470010 HCAPLUS <u>Full-text</u>

DOCUMENT NUMBER: 148:100384

TITLE: Preparation of 1,3-diphenylpropane derivatives,

particularly 2-[4-(3-oxo-3-phenylpropyl)phenoxy]-2-methylpropanoic acids and related derivatives, as PPAR agonists for treating diseases especially dyslipidemia

INVENTOR(S):
Delhomel, Jean-François; Hanf, Remy;

Caumont-Bertrand, Karine

PATENT ASSIGNEE(S): Genfit, Fr.

SOURCE: PCT Int. Appl., 97pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 2

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WO 2007147880	A1 200	71227 WO 2	007-EP56225	20070621
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GB, GD, GE,	GH, GM, GT	, HN, HR, HU,	ID, IL, IN, IS,	JP, KE, KG,
KM, KN, KP,	KR, KZ, LA	, LC, LK, LR,	LS, LT, LU, LY,	MA, MD, ME,
MG, MK, MN,	MW, MX, MY	, MZ, NA, NG,	NI, NO, NZ, OM,	PG, PH, PL,
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RW: AT, BE, BG,	CH, CY, CZ	, DE, DK, EE,	ES, FI, FR, GB,	GR, HU, IE,
IS, IT, LT,	LU, LV, MC	, MT, NL, PL,	PT, RO, SE, SI,	SK, TR, BF,
BJ, CF, CG,	CI, CM, GA	, GN, GQ, GW,	ML, MR, NE, SN,	TD, TG, BW,
GH, GM, KE,	LS, MW, MZ	, NA, SD, SL,	SZ, TZ, UG, ZM,	ZW, AM, AZ,
BY, KG, KZ,	MD, RU, TJ	, TM		
FR 2902789	A1 200	71228 FR 2	006-5540	20060621

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CA	2655	744			A1	20071227			CA 2007-2655744					20070621			
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OTHER SC	DURCE	(S):			MARI	PAT	148:	1003	84								

GΙ

$$X^{1}$$
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Title compds. I [X1 = halo, R1, G1R1; X2 = halo, R2, G2R2; X3 = R3, G3R3; X4 = halo,AΒ R4, G4R4; X5 = R5, G5R5; R1 = haloalky1; R2 = H, alky1; R3-R5 = independently H, (un) substituted alkyl; G1-G5 = independently O, S; with at least one of X3-X5 = R3, G3R3, R4, G4R4, R5, G5R5 in which G3-G5 = defined as above and R3-R5 = independently alkyl substituted with 1-2 substituents selected from CO2H and derivs., CONH2 and derivs., SO3H, SO2NH2 and derivs.; A = CR6R7, CO, C:N-OH, C:N-OR8; R6 = H, alkyl, OR8; R7 = alkyl, OH, OR8; R8 = independently alkyl substituted with an aryl or cycloalkyl group; D = CH2, CHY; Y = O- or S-heterocycle; and their stereoisomers, racemates, geometrical isomers, tautomers, salts, hydrates, solvates, solid forms and their mixts.] were prepared as PPAR activators, especially agonists, for treating dyslipidemia, diabetes type II and related diseases. Thus, reduction of 2-[2,6-dimethyl-4-[3-[4-(trifluoromethylthio)phenyl]-3-oxoprop-1-enyl]phenoxy]-2methylpropanoic acid with triethylsilane in DCM in the presence of TFA at room temperature gave the acid II (m.p. = $83-85^{\circ}$). Selected I were hPPAR α , hPPAR γ , and/or hPPA δ activators in an induced luciferase activity via hPPAR α /Gal4, hPPAR γ /Gal4, and hPPAR δ /Gal4 transactivation assay. I displayed hypolipemic properties by lowering the plasmatic cholesterol and triglycerides rates. I are useful for treating diabetes type II, dyslipidemia, pathologies associated with metabolic syndrome, cardiovascular diseases, etc.

1000336-61-7, 2-[[4-[3-(4-Chloro-2-hydroxyphenyl)-3-oxoprop-1-ΙT enyl]phenyl]thio]-2-methylpropanoic acid

RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of 1,3-diphenylpropane derivs. as PPAR activators for treating diseases especially dyslipidemia)

1000336-61-7 HCAPLUS RN

Propanoic acid, 2-[[4-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-CN yl]phenyl]thio]-2-methyl- (CA INDEX NAME)

$$HO_2C$$
 CH CH CH CH

REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 3 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:650984 HCAPLUS Full-text

DOCUMENT NUMBER: 141:190511

TITLE: Preparation of acyl aminopropanediols as PPAR, in

particular PPAR α , agonists and antioxidants for treating cerebral ischemia and related diseases

INVENTOR(S): Darteil, Raphael; Caumont, Bertrand Karine;

Najib, Jamila

PATENT ASSIGNEE(S): Genfit S. A., Fr. SOURCE: Fr. Demande, 95 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PRIOF												2003-					0030	212
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ОТНЕВ	2 20	JIIBCE	(8) •			MZD.	РΔТ	141.	1905	1 1								

OTHER SOURCE(S): MARPAT 141:190511

GΙ

Title compds. I [wherein F, G = independently O, S, NR4; F = G = NR4 never possible; AB R, R4 = independently H, (un)saturated (un)substituted alkyl; R1, R2, R3 = independently H, C(:0)R5, C(:0)(CH2)2n+1-X-R6, with a least one of R1, R2, R3 = C(:0) (CH2) 2n+1-X-R6; R5 = (un) saturated (un) substituted (C1-C25) alkyl, optionally containing a cyclic group; X = S, Se, SO, SO2; n = 0-11; R6 = (un)saturated (un)substituted (C3-C23) alkyl, optionally containing a cyclic group and/or O, S, Se, SO, SO2; with the exclusion of compds. for which FR2 = GR3 = OH; their optical and geometrical isomers, racemates, salts, hydrates and mixts.] were prepared as peroxisome proliferator-activated receptors- α (PPAR α) agonists and antioxidants for treating cerebral ischemia and related diseases. For example, II was prepared in 3 steps from 1-bromotetradecane, mercaptoacetic acid, 3-aminopropane-1,2,-diol, and palmitic acid. In an antioxidant test, selected I diminished the formation of oxidation product of LDL by AAPH by 33%. Selected I were PPAR α agonists and showed induced luciferase activity via $PPAR\alpha/Gal4$ transactivation. I are neuroprotectants useful for treating ischemia.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 4 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:650967 HCAPLUS Full-text

DOCUMENT NUMBER: 141:185113

TITLE: Therapeutic use of acyl glycerols and their nitrogen

and sulfur analogs

INVENTOR(S): Darteil, Raphael; Caumont, Bertrand Karine;

Najib, Jamila

PATENT ASSIGNEE(S): Genfit S. A., Fr. SOURCE: Fr. Demande, 144 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

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         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,
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                                                                 20040212
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                      A1 20060713 US 2005-542512
     US 20060154984
                                                                 20050718
PRIORITY APPLN. INFO.:
                                           FR 2003-1691
                                                               A 20030212
                                                               W 20040212
                                            WO 2004-FR322
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OTHER SOURCE(S): MARPAT 141:185113

The invention discloses the use of acyl glycerols and their nitrogen and sulfur analogs for the therapy and in particular in human health. The compds. of the invention have advantageous pharmacol. properties and are in particular usable for the prevention and treatment of neurodegenerative diseases.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 5 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:631313 HCAPLUS Full-text

DOCUMENT NUMBER: 141:151015

TITLE: Methods for the synthesis of nitrogen and sulphide analogs of acylglycerols and uses thereof in the

treatment of brain diseases

Darteil, Raphael; Caumont Bertrand, Karine; INVENTOR(S):

Najib, Jamila

PATENT ASSIGNEE(S): Genfit S.A., Fr. SOURCE: Fr. Demande, 86 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	ATENT				KIN	D	DATE				ICAT					ATE	
F	R 285	0650			A1		2004	0806							20030131		
	R 285						2005										
C.	A 251	4301			A1		2004	0819		CA 2	004-	2514	301		2	0040.	202
W	0 200	10692	41		A1		2004	0819		WO 2	004 - 1	FR22	9		2	0040	202
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		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,
		LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI
	RW	BW,	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑT,	BE,
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J.	P 200																202
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PRIORI	TY AP:	PLN.	INFO	.:						FR 2	003-	1144		i	A 2	0030	131
										wo 2	004-	FR22	9	Ī	W 2	0040	202

MARPAT 141:151015

The present invention relates to preparation and therapeutic use of acylglycerols and their nitrogen and sulfide analogs, in particular for the treatment of cerebral ischemia.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 6 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:19768 HCAPLUS Full-text DOCUMENT NUMBER: 140:76897

TITLE: Preparation of 1,3-diphenylprop-2-en-1-one as PPAR

agonists and as antioxidants for treating cerebral

ischemia and related diseases Najib, Jamila; Caumont Bertrand,

Karine

PATENT ASSIGNEE(S): Genfit S.A., Fr. SOURCE: Fr. Demande, 66 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 2 PATENT INFORMATION:

INVENTOR(S):

		ENT									APPLICATION NO.							
	FR	2841	900			A1		2004	0109			2002-					0020	708
		2841				В1		2007										
	CA	2490	986			A1		2004	0115		CA	2003-	2490	986		2	0030	708
	WO	2004	0052	33		A1		2004	0115		WO	2003-	FR21	27		2	0030	708
		W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB	BG,	BR,	BY,	BZ,	CA,	CH,	CN,
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC	E, EE,	ES,	FΙ,	GB,	GD,	GE,	GH,
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE	KG,	KΡ,	KR,	KΖ,	LC,	LK,	LR,
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN	I, MW,	MX,	MZ,	NI,	NO,	NZ,	OM,
			PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE	s, SG,	SK,	SL,	SY,	ΤJ,	TM,	TN,
							•					I, YU,						
		RW:		•	•	•	•	•	•			TZ,		•			•	•
												СН,						
												NL,						
												, GW,						
		2003				A1						2003-						
		2003										2003-						
		1525				A1					EΡ	2003-	7627	49		2	0030	708
	EΡ	1525						2007										
		R:										I, IT,						PT,
			•									, TR,						
		1668				А		2005	0914		CN	2003-	8163	66		2	0030	
		2005	5323	85		T						2004-					0030	
		3657	03			T						2003-					0030	
		5380	51			A		2007				2003-					0030	
		2287				Т3		2007				2003-					0030	
		2004						2005			NO	2004-	530I			2	0041	
		2005				A		2005			MX	2005-	427			2	0050	
		2005				A		2007			ZΑ	2005- 2005-	1082			2	0050	
		2005				A1		2005			US	2005-	5200	19		2	0050	
DDTO		2007				A1		2007	UZU8			2006-					0060	
PRIOF	< T T, 7	APP.	ыN•.	TNE.O	.:							2002-					0020	
												2003-					0030	
											US	2005-	5200	19		AZ Z	0050	422

OTHER SOURCE(S): MARPAT 140:76897

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^{*} STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

Title compds. I [wherein X1 = halo, R1, G1R1; X2 = H, thionitroso, OH, AΒ alkylcarbonyloxy, alkyloxy, SH, alkylthio, alkylcarbonylthio or X2 = O or S that forms a 2-phenyl-4H-1-benzopyran-4-one with the carbon-3 of the propene chain; X3 = R3, G3R3; X4 = halo, thionitroso, R4, G4R4; X5 = R5, G5R5; X6 = O, NH and derivs.; R1, R3, R4, R5 = independently H, (un)substituted alkyl; G1, G3, G4, G5 = independently O or S; with at least one of X1, X3, X4, or X5 of formula GR and one of the R1, R3, R4, or R5 is a substituted radical, and that radical form a cycle, or is associated with a group G; their optical and geometrical isomers, racemates, tautomers, salts, hydrates and mixts.; with the exclusion of certain compds.] were prepared as peroxisome proliferator-activated receptors- α (PPAR α) agonists and as antioxidants for treating cerebral ischemia and related diseases. For example, II was prepared by mixed-Aldol condensation of ketone III with 4-hydroxy-3,5ditertbutylbenzaldehyde in the presence of ethanol/HCl. In an antioxidant test, selected I diminished the formation of oxidation product of LDL by AAPH by 33%. Selected I were PPAR α agonists and showed induced luciferase activity via PPAR α /Gal4 transactivation. I are neuroprotectants useful for treating ischemia.

REFERENCE COUNT: 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L23 ANSWER 7 OF 7 HCAPLUS COPYRIGHT 2009 ACS on STN ACCESSION NUMBER: 2004:19750 HCAPLUS Full-text

DOCUMENT NUMBER: 140:76896

TITLE: Composition based on substituted

1,3-diphenylprop-en-1-one derivatives, preparation and

use as PPARlpha agonists, antioxidants as well as

antiinflammatory agents

INVENTOR(S): Najib, Jamila; Caumont Bertrand,

Karine

PATENT ASSIGNEE(S): Genfit S.A., Fr. SOURCE: Fr. Demande, 66 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT NO.	KIND DATE	APPLIC	CATION NO.	DATE
FR 2841784	A1 2004	:0109 FR 200)2-8570	20020708
FR 2841784	B1 2007	0302		
CA 2490993	A1 2004	:0115 CA 200)3-2490993	20030708
WO 2004005243	A2 2004	:0115 WO 200)3-FR2128	20030708
WO 2004005243	A3 2004	0422		
W: AE, AG, AL,	AM, AT, AU,	AZ, BA, BB, B	BG, BR, BY, BZ,	CA, CH, CN,
CO, CR, CU,	CZ, DE, DK,	DM, DZ, EC, E	EE, ES, FI, GB,	GD, GE, GH,
GM, HR, HU,	ID, IL, IN,	IS, JP, KE, K	KG, KP, KR, KZ,	LC, LK, LR,
LS, LT, LU,	LV, MA, MD,	MG, MK, MN, M	MW, MX, MZ, NI,	NO, NZ, OM,
PG, PH, PL,	PT, RO, RU,	SC, SD, SE, S	SG, SK, SL, SY,	TJ, TM, TN,
TR, TT, TZ,	UA, UG, US,	UZ, VC, VN, Y	YU, ZA, ZM, ZW	
RW: GH, GM, KE,	LS, MW, MZ,	SD, SL, SZ, I	ZZ, UG, ZM, ZW,	AM, AZ, BY,
KG, KZ, MD,	RU, TJ, TM,	AT, BE, BG, C	CH, CY, CZ, DE,	DK, EE, ES,
FI, FR, GB,	GR, HU, IE,	IT, LU, MC, N	NL, PT, RO, SE,	SI, SK, TR,
BF, BJ, CF,	CG, CI, CM,	GA, GN, GQ, G	GW, ML, MR, NE,	SN, TD, TG
AU 2003264699	A1 2004	.0123 AU 200	3-264699	20030708
EP 1519908	A2 2005	0406 EP 200	3-762750	20030708
EP 1519908	B1 2007	0613		
R: AT, BE, CH,	DE, DK, ES,	FR, GB, GR, I	T, LI, LU, NL,	SE, MC, PT,
IE, SI, LT,	LV, FI, RO,	MK, CY, AL, T	CR, BG, CZ, EE,	HU, SK
BR 2003012399	A 2005	0412 BR 200	3-12399	20030708
CN 1688532	A 2005	1026 CN 200	3-816351	20030708

Serial#: 10/520,078 Τ 20051027 JP 2004-518891 JP 2005532386 20030708 AT 364588 20070715 AT 2003-762750 Τ 20030708 NZ 538052 20070928 NZ 2003-538052 A 20030708 20071216 ES 2003-762750 20041227 NO 2004-5082 Т3 ES 2287529 20030708 A NO 2004005082 20041122 MX 2005000425 20050722 MX 2005-425 Α 20050107 ZA 2005001081 Α 20070425 ZA 2005-1081 20050207 US 2005-520078 20050404 FR 2002-8570 A 20020708 WO 2003-FR2128 W 20030708 US 20050171149 A1 20050804 US 2005-520078 PRIORITY APPLN. INFO.:

OTHER SOURCE(S): MARPAT 140:76896

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- * STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY AVAILABLE VIA OFFLINE PRINT *
- Title compds. I [wherein X1 = halo, R1, G1R1; X2 = H, thionitroso, OH, AB alkylcarbonyloxy, alkyloxy, SH, alkylthio, alkylcarbonylthio or X2 = O or S that forms a 2-phenyl-4H-1-benzopyran-4-one with the carbon-3 of the propene chain; X3 =R3, G3R3; X4 = halo, thionitroso, R4, G4R4; X5 = R5, G5R5; X6 = O, NH and derivs.; R1, R3, R4, R5 = independently H, (un) substituted alkyl; G1, G3, G4, G5 = independently O or S; with at least one of X1, X3, X4, or X5 of formula GR and one of the R1, R3, R4, or R5 is a substituted radical, and that radical form a cycle, or is associated with a group G; their optical and geometrical isomers, racemates, tautomers, salts, hydrates and mixts.; with the exclusion of certain compds.] were prepared as peroxisome proliferator-activated receptors- α (PPAR α) agonists and as antioxidants for treating cerebral ischemia and related diseases. For example, II was prepared by mixed-Aldol condensation of ketone III with 4-hydroxy-3,5ditertbutylbenzaldehyde in the presence of ethanol/HCl. In an antioxidant test, selected I (10-3 M) diminished the formation of oxidation product of LDL by AAPH by Selected I were PPARlpha agonists, showing induced luciferase activity via $PPAR\alpha/Gal4$ transactivation with a factor of induction ranging from 10 to 60, 5-50 and 3-35 at 100 μM , 30 μM , and 10 μM resp. I and their compns. are useful for treating cardiovascular diseases, syndrome X, restenosis, diabetes, obesity, hypertension, inflammatory diseases, cancers or neoplasms (benign or malignant tumors), neurodegenerative diseases, dermatol. and the disorders related to the oxydative stress, for preventing and treating aging, and in particular cutaneous

IT 639864-16-7P 639864-17-8P 639864-18-9P 639864-19-0P 639864-20-3P 639864-21-4P

639864-22-5P 639864-23-6P 639864-30-5P

639864-31-6P 639864-38-3P 639864-39-4P

639864-40-7P 639864-41-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(PPAR α agonist; preparation of diphenylpropenones as PPAR agonists for treating ischemia)

RN 639864-16-7 HCAPLUS

CN Propanoic acid, 2-[3-(1,1-dimethylethyl)-2-hydroxy-5-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl- (CA INDEX NAME)

$$\begin{array}{c} \text{Me} \\ \text{Ho}_2\text{C} \\ \text{OH} \end{array} \begin{array}{c} \text{OH} \\ \text{Bu} \\ \text{OH} \end{array}$$

RN 639864-17-8 HCAPLUS

CN Propanoic acid, 2-[3-(1,1-dimethylethyl)-2-hydroxy-5-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-18-9 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-3-(1,1-dimethylethyl)-2-hydroxyphenoxy]-2-methyl- (CA INDEX NAME)

RN 639864-19-0 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-3-(1,1-dimethylethyl)-2-hydroxyphenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-20-3 HCAPLUS

CN Benzeneacetic acid, $3-(1,1-dimethylethyl)-2-hydroxy-5-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-<math>\alpha$, α -dimethyl- (CA INDEX NAME)

RN 639864-21-4 HCAPLUS

CN Benzeneacetic acid, $3-(1,1-\text{dimethylethyl})-2-\text{hydroxy}-5-[3-(2-\text{hydroxyphenyl})-3-\text{oxo}-1-\text{propen}-1-\text{yl}]-\alpha,\alpha-\text{dimethyl}-$, 1-methylethyl ester (CA INDEX NAME)

RN 639864-22-5 HCAPLUS

CN Benzeneacetic acid, $5-[3-(4-\text{chloro}-2-\text{hydroxyphenyl})-3-\text{oxo}-1-\text{propen}-1-\text{yl}]-3-(1,1-\text{dimethylethyl})-2-\text{hydroxy}-\alpha,\alpha-\text{dimethyl}-$ (CA INDEX NAME)

RN 639864-23-6 HCAPLUS

CN Benzeneacetic acid, $5-[3-(4-\text{chloro}-2-\text{hydroxyphenyl})-3-\text{oxo}-1-\text{propen}-1-\text{yl}]-3-(1,1-\text{dimethylethyl})-2-\text{hydroxy}-\alpha,\alpha-\text{dimethyl}-$, 1-methylethyl ester (CA INDEX NAME)

RN 639864-30-5 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-2,3-dihydroxyphenoxy]-2-methyl- (CA INDEX NAME)

$$\begin{array}{c} \text{C1} \\ \text{OH} \\ \text{OH} \end{array}$$

RN 639864-31-6 HCAPLUS

CN Propanoic acid, 2-[5-[3-(4-chloro-2-hydroxyphenyl)-3-oxo-1-propen-1-yl]-2,3-dihydroxyphenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-38-3 HCAPLUS

CN Propanoic acid, 2-[3-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl- (CA INDEX NAME)

RN 639864-39-4 HCAPLUS

CN Propanoic acid, 2-[3-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenoxy]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

RN 639864-40-7 HCAPLUS

CN Propanoic acid, 2-[[4-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenyl]thio]-2-methyl- (CA INDEX NAME)

RN 639864-41-8 HCAPLUS

CN Propanoic acid, 2-[[4-[3-(2-hydroxyphenyl)-3-oxo-1-propen-1-yl]phenyl]thio]-2-methyl-, 1-methylethyl ester (CA INDEX NAME)

REFERENCE COUNT:

29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

Serial#: 10/520,078 SEARCH HISTORY

		STRY' ENTER		39:25 ON	30 JUN 2009
L1 L2	47768	STR SEA SSS FU	L L1		
		ACT ZARREG	21SB/A		
L3		STR			
L4	(47768))SEA SSS FU	L L3		
L5		STR			
L6	2679	SEA SUB=L4	SSS FUL	L5	
	FILE 'HCAP	LUS' ENTERE	D AT 11:4	4:16 ON	30 JUN 2009
L7		SEA SPE=ON			
L8	191239				(DIABETES/CT OR "DIABETES INSIPIDUS
					/CT) OR ?DIABET?/BI
L9	74644	SEA SPE=ON LER?/BI	ABB=ON	PLU=ON	ATHEROSCLEROSIS+OLD/CT OR ?ATHEROSC
L10	187726		ABB=ON	PLH=ON	OBESITY+NT/CT OR ?OBESIT?/BI OR
што	10,,20				WT) (5A) (LOSS OR GAIN OR REDUCTION
		OR MANAGEMI			, (, (
L11	3805	SEA SPE=ON	ABB=ON	PLU=ON	L8 AND L9 AND L10
L12	5	SEA SPE=ON	ABB=ON	PLU=ON	L7 AND L11
L13	24	SEA SPE=ON	ABB=ON	PLU=ON	L7(L)L8
L14	3	SEA SPE=ON	ABB=ON	PLU=ON	L7(L)L9
L15	8	SEA SPE=ON	ABB=ON	PLU=ON	L7(L)L10
L16	31	SEA SPE=ON	ABB=ON	PLU=ON	L13 OR L14 OR L15
L17	30	SEA SPE=ON	ABB=ON	PLU=ON	L16 NOT L12
L18	16	SEA SPE=ON	ABB=ON	PLU=ON	L17 AND (PRY<=2003 OR AY<=2003 OR
		PY<=2003 O			
L19					NAJIB J?/AU
L20	90				CAUMONT-BERTRAND K?/AU OR CAUMONT
		K?/AU OR BI		•	
L21					L19 AND L20
L22					(L12 OR L18) AND (L19 OR L20)
L23	7	SEA SPE=ON	ABB=ON	PLU=ON	L21 OR L22